

This Page Is Inserted by IFW Operations
and is not a part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images may include (but are not limited to):

- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

IMAGES ARE BEST AVAILABLE COPY.

**As rescanning documents *will not* correct images,
please do not report the images to the
Image Problems Mailbox.**

This Page Blank (uspto)

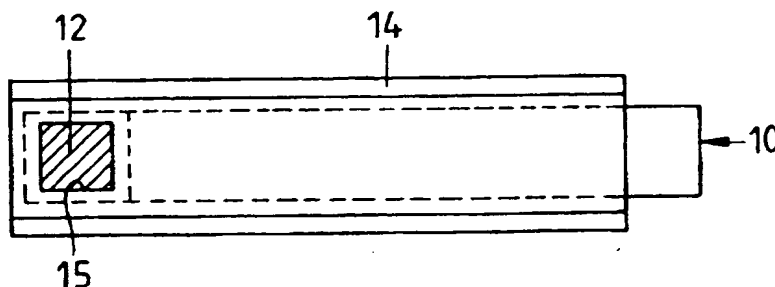


INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁶: G01N 1/00, 1/28, 30/90, 33/53	A1	(11) International Publication Number: WO 96/28715 (43) International Publication Date: 19 September 1996 (19.09.96)
(21) International Application Number: PCT/AU96/00135 (22) International Filing Date: 13 March 1996 (13.03.96) (30) Priority Data: PN 1737 14 March 1995 (14.03.95) AU PN 5597 25 September 1995 (25.09.95) AU (71)(72) Applicant and Inventor: CHANDLER, Howard, Milne [AU/US]; 76A Princes Point Road, Yarmouth, ME 04096 (US). (74) Agents: SLATTERY, John, Michael et al.; Davies Collison Cave, 1 Little Collins Street, Melbourne, VIC 3000 (AU).		(81) Designated States: AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, ARIPO patent (KE, LS, MW, SD, SZ, UG), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG). Published <i>With international search report.</i>

(54) Title: SAMPLE COLLECTION DEVICE**(57) Abstract**

A sample collection device (10), particularly for use in collection of faecal samples for occult blood detection, comprises a collection member having at least one absorbent matrix (12) thereon, and a protective cover member (14) for the collection member, the protective cover member being adapted to receive and shield at least the or each absorbent matrix of the collection member when the collection member is assembled therewith.



FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AM	Armenia	GB	United Kingdom	MW	Malawi
AT	Austria	GE	Georgia	MX	Mexico
AU	Australia	GN	Guinea	NE	Niger
BB	Barbados	GR	Greece	NL	Netherlands
BE	Belgium	HU	Hungary	NO	Norway
BF	Burkina Faso	IE	Ireland	NZ	New Zealand
BG	Bulgaria	IT	Italy	PL	Poland
BJ	Benin	JP	Japan	PT	Portugal
BR	Brazil	KE	Kenya	RO	Romania
BY	Belarus	KG	Kyrgyzstan	RU	Russian Federation
CA	Canada	KP	Democratic People's Republic of Korea	SD	Sudan
CF	Central African Republic	KR	Republic of Korea	SE	Sweden
CG	Congo	KZ	Kazakhstan	SG	Singapore
CH	Switzerland	LI	Liechtenstein	SI	Slovenia
CI	Côte d'Ivoire	LK	Sri Lanka	SK	Slovakia
CM	Cameroon	LR	Liberia	SN	Senegal
CN	China	LT	Lithuania	SZ	Swaziland
CS	Czechoslovakia	LU	Luxembourg	TD	Chad
CZ	Czech Republic	LV	Latvia	TG	Togo
DE	Germany	MC	Monaco	TJ	Tajikistan
DK	Denmark	MD	Republic of Moldova	TT	Trinidad and Tobago
EE	Estonia	MG	Madagascar	UA	Ukraine
ES	Spain	ML	Mali	UG	Uganda
FI	Finland	MN	Mongolia	US	United States of America
FR	France	MR	Mauritania	UZ	Uzbekistan
GA	Gabon			VN	Viet Nam

SAMPLE COLLECTION DEVICE

FIELD OF THE INVENTION

5 This invention relates to a device for collecting a sample, such as a biological sample, for subsequent use in the detection of an analyte in the sample. In one particular embodiment, this invention relates to a device for collecting a faecal sample for the purposes of subsequent occult blood detection in the sample. It is to be understood however that the device of this invention
10 may be used in the collection of other biological samples such as blood, urine, saliva samples and the like, as well as in the collection of non-biological samples such as water samples for detection of pollutants and the like.

 The present invention also extends to an assay device based on this
15 sample collection device.

BACKGROUND OF THE INVENTION

 A well known and widely-used clinical reagent for the detection of occult blood in a sample, particularly a faecal sample, is guaiac (also known as gum
20 guaiac or resin guaiac). When used in association with an appropriate developer solution, guaiac provides a colorimetric assay system for detecting haemoglobin in the sample. Such tests are commercially available, for example, Hemoccult II and Hemoccult II Sensa (SmithKline Diagnostics, San Jose, California, USA).

25

 Prior Australian Patent Application No. 21852/92 (International Patent Application No. PCT/US92/04425) notes that among the many analytical systems used for detection and/or determination of analytes, particularly analytes of biological interest, are chromatographic assay systems. Among the analytes of
30 biological interest frequently assayed with such systems are:

1. hormones, such as human chorionic gonadotropin (hCG), frequently assayed as a marker of human pregnancy;

- 2 -

2. antigens, particularly antigens specific to bacterial, viral, and protozoan pathogens, such as *Streptococcus*, hepatitis virus, and *Giardia*;
3. antibodies, particularly antibodies induced as a result of infection with pathogens, such as antibody to the bacterium *Helicobacter pylori* and to human immunodeficiency virus (HIV);
4. other proteins, such as haemoglobin, frequently assayed in determinations of faecal occult blood, an early indicator of gastrointestinal disorders such as colon cancer;
5. enzymes, such as aspartate aminotransferase, lactate dehydrogenase, alkaline phosphatase, and glutamate dehydrogenase, frequently assayed as indicators of physiological function and tissue damage;
6. drugs, both therapeutic drugs, such as antibiotics, tranquillisers and anticonvulsants, and illegal drugs of abuse, such as cocaine, heroin, and marijuana; and
7. vitamins.

Such chromatographic systems are frequently used by physicians and medical technicians for rapid in-office diagnosis and therapeutic monitoring of a variety of conditions and disorders. They are also increasingly used by patients themselves for at-home monitoring of such conditions and disorders.

Among the most important of such chromatographic systems are the "thin layer" systems in which a solvent moves as a solvent front across a thin, flat absorbent medium. Among the most important of tests that can be performed with such thin layer systems are immunoassays, which depend on the specific interaction between an antigen or hapten and a corresponding antibody. The use of immunoassays as a means of testing for the presence and/or amount of clinically important molecules has been known for some time.

Chromatographic techniques used in conjunction with immunoassays include a procedure known as immunochromatography. In general, this technique uses a disclosing reagent or particle that has been linked to an

antibody to the analyte to be assayed, forming a conjugate. This conjugate is then mixed with a specimen and, if the analyte to be assayed is present in the specimen, the disclosing reagent-linked antibodies bind to the analyte to be assayed, thereby giving an indication that the analyte to be assayed is present.

- 5 The disclosing reagent or particle can be identifiable by colour, magnetic properties, radioactivity, specific reactivity with another molecule, or another physical or chemical property. The specific reactions that are employed vary with the nature of the analyte being assayed and the sample to be tested.

- 10 Although useful, currently available chromatographic techniques using test strips have a number of drawbacks. Many samples, such as faecal samples, contain particulate matter that can clog the pores of the chromatographic medium, greatly hindering the immunochromatographic process. Other samples, such as blood, contain cells and coloured components that make it difficult to
15 read the test. Even if the sample does not create interference, it is frequently difficult with existing chromatographic test devices to apply the sample to the chromatographic medium so that the solvent front moves uniformly through the chromatographic medium to ensure that the sample reaches the area where binding is to occur in a uniform, straight-line manner.

20

- Sample preparation and waste generation are responsible for other problems with currently available devices and techniques for immunochromatography. The increased prevalence of diseases spread by infected blood and blood fractions, such as AIDS and hepatitis, has exacerbated
25 these problems. It is rarely possible to apply a sample (such as faeces) or a sampling device (such as a throat swab) directly to the chromatographic medium. Several extraction and pretreatment reactions are usually required before the sample can be applied to the chromatographic medium. These reactions are typically carried out by the physician or technician performing the
30 test in several small vessels, such as test tubes or microfuge tubes, requiring the use of transfer devices, such as pipettes. Each of these devices is then contaminated and must be disposed of as waste using special precautions so

- 4 -

that workers or people who may inadvertently come into contact with the waste do not become contaminated.

The present invention is particularly, but not exclusively, directed to collection of faecal samples for occult blood detection, for example in screening for colorectal cancer. As previously described, guaiac testing provides a colorimetric assay system for detection of haemoglobin in a sample, however because of the large number of false positives obtained in guaiac testing, in screening programs the use of two or three guaiac tests has been recommended, confirmed when positive by an immunological test for human haemoglobin (Favennic L., Kapel N., Meillet D., Chochillon C. and Gobert J.G., *Annales de Biologie Clinique*, 50(5):311-3, 1992). More recently, a combination of guaiac and immunological testing has been suggested (Allison, J.E., Tekawa, I.S., Ransom, L.J. and Adrian, L.L. *N. Engl. J. Med.*, 334:155-9, 1996).

15

It is an object of the present invention to provide a sample collection device which is simple and economic to manufacture and which enables subsequent detection and/or determination of analyte in the sample to be readily carried out, for example using an immunochromatographic or other immunodiagnostic procedure. It is a particular object of the present invention to provide a sample collection device which is suitable for use in testing for faecal occult blood in a combination of guaiac and immunological testing.

20

SUMMARY OF THE INVENTION

In accordance with the present invention, there is provided a sample collection device comprising

25

- (a) a collection member having at least one absorbent matrix thereon; and
- (b) a protective cover member for said collection member, said protective cover member being adapted to receive and shield at least the absorbent matrix of said collection member when the collection member is assembled therewith.

30

- 5 -

In another aspect, the invention also provides an assay device for detection and/or determination of an analyte in a sample, comprising:

- (a) a collection member having at least one absorbent matrix thereon adapted to receive said sample,
- 5 (b) a protective cover member for said collection member, said protective cover member being adapted to receive and shield at least the or each absorbent matrix of said collection member when the collection member is assembled therewith, and
- (c) means to detect and/or determine said analyte in said sample.

10

Throughout this specification and the claims which follow, unless the context requires otherwise, the word "comprise", or variations such as "comprises" or "comprising", will be understood to imply the inclusion of a stated integer or group of integers but not the exclusion of any other integer or group

15 of integers.

DETAILED DESCRIPTION OF THE INVENTION

Preferably, the collection member is an elongate member in the form of a dipstick, test strip or the like. Such an elongate collection member may

20 comprise an elongate rigid or semi-rigid carrier or backing having the absorbent matrix affixed to the carrier or backing at or towards one end thereof. Typically, the elongate collection member may comprise a dipstick in the form of a flat, generally rectangular carrier strip, with the or each absorbent matrix being located at one end of the strip, and the opposite end of the strip providing

25 means for handling the strip.

Suitable materials for use as a carrier or backing in such an elongate collection member include, for example, plastics materials such as polycarbonate, polyethylene, Mylar, vinyl, cellophane and polystyrene, as well

30 as water-proofed or water-resistant cardboard or similar materials. In some embodiments of this invention, the carrier or backing is preferably made from a transparent or translucent material.

The or each absorbent matrix which is located on the collection member, preferably at or towards one end of a carrier strip, may comprise any suitable absorbent material. A particularly preferred material is absorbent paper such as filter paper, however any other absorbent material may also be used including, by way of example, cellulose, nitrocellulose, nylon, rayon, glass fibre, sintered glass, fleeces or non-woven or porous synthetic materials.

In one particular embodiment, the absorbent matrix on one or both sides of the collection member may comprise a sample collection matrix in operable contact with an absorbent reagent matrix or capillary means, so that reagent added to the absorbent reagent matrix or capillary means may flow along the collection member and into the sample collection matrix. If desired, at least the region of the junction between the sample collection matrix and the absorbent reagent matrix or capillary means may be covered with an impervious protective layer to prevent contamination of the absorbent reagent matrix or capillary means when a sample such as a faecal sample is applied to the sample collection matrix.

The absorbent matrix or matrices may be affixed to a carrier strip by means of an adhesive. Suitable adhesives are well-known in the art.

In order to assist in the collection of sample material, particularly faecal material, it is preferred that some means be provided for removal of excess sample material, particularly particulate material, after it has been applied to the absorbent matrix on the collection member. One such approach is to simply wipe the collection member with an absorbent material. Alternatively, a wiper sleeve of plastic backed absorbent paper or the like may be supplied on the collection member to clean away the excess material. In another embodiment, the collection member may be provided with an optionally removable protective cover layer or sheath which may be provided with a window or aperture whereby only the absorbent matrix or a portion thereof is exposed, so that after application of faecal material to the exposed portion of the absorbent matrix, the

collection member may be removed from its protective sheath or the protective cover layer removed leaving excess faecal material to be disposed of with the cover layer or sheath, and the collection member may then be used in further testing as described herein.

5

Preferably, the protective cover member of the sample collection device of this invention is in the form of a sheath or housing having an internal recess which is adapted to receive the collection member and shield the absorbent matrix or matrices. In a particularly preferred embodiment the internal recess
10 in the sheath or housing receives substantially the entire length of an elongate collection member.

The protective cover member is preferably made of a water-impervious, non-absorbent material. Suitable materials include plastics materials such as
15 polycarbonate, and water-proofed or water-resistant cardboard or similar materials, as well as laminated materials such as plasticised cardboard or metal foil, for example plasticised aluminium foil. In one simple and inexpensive embodiment, the protective cover member may be made from a single flat piece of suitable material which is folded along one fold line to give two opposed
20 panels which are then affixed at the edges, either with a suitable adhesive or by welding or fusing or the like, to form a sheath or housing with a suitable internal recess. In other embodiments, the initial piece of material may be folded along two or more fold lines to provide a sheath or housing as described above as a first opposable component together with one or more additional opposable
25 components which are hingedly joined to the first opposable component and can be brought into opposition with the first opposable component during the test procedure.

In accordance with this invention, the protective cover member receives
30 and shields at least the absorbent matrix or matrices of the collection member when assembled therewith. Thus, when a sample such as a faecal sample or other sample of biological interest is applied to the absorbent matrix or matrices

- 8 -

and the collection member is assembled within the protective cover member, persons who handle the assembled device are protected against exposure to the sample and any bacterial, viral or protozoan pathogens which may be present in the sample.

5

In particular embodiments of the invention, the protective cover member may be provided with at least one window or aperture through a wall or panel thereof, the or each window or aperture being located so that it corresponds with at least a portion of an absorbent matrix when the collection member is assembled within the protective cover member. In these embodiments, the window or aperture enables test reagent(s) to be applied to a sample on the absorbent matrix during detection and/or determination of analyte(s) in the sample, without the need for removal of the collection device from the protective cover member. Such a window or aperture is preferably covered with a removable, and optionally resealable, sealing layer so that the window or aperture is covered by the sealing layer during collection and handling of the sample and then exposed so that test reagent(s) can be added to the sample when the test analysis is to be performed. Where the sealing layer is resealable, this layer can be reapplied to cover the window or aperture after the addition of the test reagent(s).

In further embodiments, one or more additional windows or apertures may be provided in the protective cover member. By way of example, such an additional window or aperture may be located so that, in conjunction with the use of a transparent or translucent carrier for the collection member, a reading window or aperture is provided for reading the results of a chromatographic test analysis through the transparent or translucent carrier. Once again, such additional windows or apertures may be covered with a sealing layer, which may be removable and resealable. Preferably, the sealing layer is transparent if the window or aperture is to be used to read test results.

The protective cover member may also include test reagent(s) sealed therein with a frangible seal, so that the test reagent(s) are released when the frangible seal is broken or perforated, for example on insertion of the collection member into the cover member.

5

In one embodiment of the sample collection device of this invention for use in faecal occult blood testing, at least a portion of the or each absorbent matrix on the collection member is impregnated with guaiac. In a particularly preferred embodiment for this use, the collection member is provided with two or more absorbent matrices and one of these is impregnated with guaiac, the other or others providing a sample taken simultaneously which is available for immunological testing.

Alternatively, or additionally, the sample collection device may further comprise a test strip located or adapted to be located on the collection member or on or in the protective cover member. Suitable test strips include a chromatographic medium, for example, immunochromatographic strips.

As previously described, the present invention also extends to an assay device which comprises, in addition to a collection member and a protective cover member as described above, means to detect and/or determine an analyte in the collected sample. Such means may include guaiac impregnated in the or each absorbent matrix, or alternatively or additionally a test strip (such as an immunochromatographic strip) located or adapted to be located on the collection member or on or in the protective cover member.

Various features of a number of embodiments of the present invention are illustrated by way of example in the accompanying drawings which are included by way of illustration, not limitation of this invention.

30

In the accompanying drawings:

- 10 -

Figure 1A is a plan view of a dipstick collection device in accordance with one embodiment of the present invention, and

Figure 1B is a side elevation of the device of Figure 1A.

5

Figure 2 is a plan view of a dipstick collection device in accordance with Figure 1A, provided with a wiper sleeve as described below.

Figure 3A is a plan view of another embodiment of a dipstick collection device in accordance with the invention, and

10

Figure 3B is an exploded side elevation of the device of Figure 3A.

Figure 4A is a plan view of a further embodiment of a dipstick collection device in accordance with this invention, and

15

Figure 4B is an exploded side elevation of the device of Figure 4A.

Figure 5A is a plan view from one side of an assembled sample collection device in accordance with a further embodiment of this invention which is particularly intended for use in the guaiac test for faecal occult blood, and

20

Figure 5B is a plan view of the reverse side of the sample collection device of Figure 5A.

25

Figures 6A and 6B are plan views from opposite sides of a sample collection device in accordance with another embodiment of this invention which is particularly intended for use in immunodiagnostic testing for faecal occult blood, and

30

Figure 6C is a plan view of the device of Figures 6A and 6B in use in immunodiagnostic testing.

Figure 7A is a diagrammatic illustration of the folding of a single piece of material such as cardboard to give a two-panel protective cover member for another embodiment of a sample collection device in accordance with this invention,

5

Figure 7B is a plan view of the protective cover member of Figure 7A in "open" position prior to insertion of the dipstick collection device,

Figure 7C is a plan view of the assembled sample collection device of Figures 7A and 7B for use in immunochromatographic testing; and

10

Figure 7D shows diagrammatically the liquid flow paths between test strip and dipstick when a test is run using the device of Figures 7A and 7B.

Figure 8A is a plan view of a modified dipstick collection device of this invention,

15

Figure 8B is an exploded side elevation of the device of Figure 8A, and

Figure 8C is a plan view of a sheath for the dipstick collection device of Figure 8A.

20

Figures 9A and 9B illustrate diagrammatically a further modification of the sample collection device of this invention.

25

Figure 10 shows diagrammatically an assembly of a dipstick collection device in a disposable protective sheath.

Figures 11A to 11D are plan views from opposite sides of a further embodiment of a sample collection device in accordance with this invention which is particularly intended for use in the guaiac test for faecal occult blood.

30

- 12 -

The dipstick collection device (10) shown in Figures 1A and 1B comprises a semi-rigid elongate backing (11) having an absorbent matrix (12) adhered to one side of one end of the backing (11). Alternatively, an absorbent matrix (12) may be located on each side of one end of the backing (11). As shown in Figure 2, in use a wiper sleeve (13) may be used to wipe the dipstick (10) as the dipstick is withdrawn from the sleeve. The sleeve may be provided in the form of a preformed sleeve which may already be in place around the elongate semi-rigid backing (11).

Figure 10 shows an alternative embodiment in which the dipstick collection device (10) is provided within a disposable protective sheath (14) which may be made from paper, cardboard, plastic, foil or any other suitable material. The sheath (14) is provided with at least one opening (15) in registry with the absorbent matrix (12) of the device (10). Alternatively, if an absorbent matrix is provided on each side of the device (10), the sheath may be provided with an opening on each side thereof so that sample is collected on both sides of the dipstick device (10), thereby enabling two tests to be performed simultaneously or sequentially on the same sample. For example, both guaiac and immunochromatographic tests for faecal occult blood may be performed on the same sample. In use, the sheathed dipstick device is inserted into a faecal specimen so that a sample is applied, via the opening(s) in the base of the sheath, to the appropriate absorbent matrix region on one or both sides of the dipstick device. If desired, the openings may be covered with a screening or filter material to prevent the collection of particulate material. The sheath (14) is held while the dipstick device is withdrawn from the specimen, after which the sheath (14) is discarded with the excess faecal specimen. An alternative sheath (14) could be constructed of a biodegradable material such as paper or cardboard. The dipstick device (10) would be placed and held within the closed sheath (14) while the sample was taken. On withdrawal of the dipstick device, the sheath (14) would fall open for discarding, e.g. flushing, with the faeces.

- 13 -

Figures 3A and 3B illustrate another embodiment of a dipstick collection device (30) in which the absorbent matrix such as guaiac test paper (32) is adhered to one side and at one end of a transparent elongate backing (31) by means of a layer of double sided adhesive (33). A covering member (34) is laminated to the semi-rigid backing (31) in such a way that a portion of the absorbent guaiac test paper (32) is covered by the covering membrane (34), whilst the remainder of the guaiac test paper (32) is exposed for application of a test sample thereto.

A further embodiment of a dipstick collection device is shown in Figures 4A and 4B where the dipstick collection device 40 comprises a semi-rigid backing (41) having an absorbent collection matrix (42) adhered at one end thereof by means of double sided adhesive (43). Optionally, a filter screen (44) is provided over the absorbent collection matrix (42), and a peel-off impermeable cover (45) is provided over at least the end of the backing (41) where the absorbent collection matrix (42) is located. A gap (46) is provided in the impermeable cover (45) coincident with the absorbent collection matrix (42) (and filter screen (44) where provided) so that a sample such as a faecal sample applied to the impermeable cover can pass through the gap (46) and be collected by the absorbent collection matrix (42). The peel-off impermeable cover (45) may then be removed and discarded after sample collection so as to remove excess sample material from the dipstick collection device.

Figures 5A and 5B show an assembled sample collection device (50) in accordance with the present invention which is particularly intended for use in the guaiac test for faecal occult blood. Device (50) comprises a dipstick collection device (51) which may be a device of the type described in any one of Figures 1 to 4 or 10 above, together with a protective cover member or housing (52) which is adapted to receive and shield the absorbent matrix of the dipstick collection device when the collection device is assembled with the protective cover member. As shown in Figure 5A, cover member (52) is provided with a window, aperture or perforations (53) through one side wall or

panel thereof, the window, aperture or perforations being located to correspond with at least a portion of the absorbent matrix of the dipstick collection device (51) when the device (51) is assembled within the protective cover member or housing (52). As shown in Figure 5B, the protective cover member or housing
5 (52) may also be provided with a reading window (54) on the other side of the housing. The use of the device of Figures 5A and 5B in guaiac testing is described in detail below.

Figures 11A to 11D show another embodiment of a sample collection
10 device for use in guaiac testing. Figures 11A and 11B show the device (110) from each side in "open" condition prior to faecal sample collection, while Figures 11C and 11D show the device (110) from each side in "closed" condition after sample collection. In current commercial guaiac tests for faecal occult blood, the faecal specimen is smeared on one side of the guaiac paper and the
15 test reagent for development is applied to the reverse side of the guaiac paper. A similar approach is available in the embodiment of Figures 11A to 11D, which in addition allows two tests to be performed on the same faecal specimen.

Figure 11A shows the "open" dipstick device (110) with the windows (111)
20 side uppermost. Figure 11B shows the opposite side of the "open" dipstick device (110) with guaiac paper (112) located near the fold in the elongate backing (113) of the device. Figure 11C shows the dipstick device (110) folded ready for faecal collection, while Figure 11D shows the dipstick device (110) folded in the reverse direction, with the faecal specimen hidden and the clean
25 side of the guaiac paper exposed for addition of the test reagent(s).

The dipstick device (110), supplied folded as shown in Figure 11C, is inserted or wiped with faecal specimen so that the guaiac paper exposed in the area of windows (111) both sides of the device (110) is contacted with
30 specimen. The dipstick device is then cleaned (e.g. wiped with toilet tissue or a protective sheath/cover peeled off and discarded) so that faecal material remains only in the window area. The patient then folds the dipstick in the

- 15 -

reverse direction as shown in Figure 11D, inserts it into its protective cover member as described herein and sends it to the laboratory or health professional for development in the usual manner. If desired, a peelable or tearable seal may be provided along the edges of the device (110) in its folded condition as
5 shown in Figure 11C to prevent entry of faecal material to the guaiac paper (112) except through the windows (111).

Figures 6A and 6B illustrate a further embodiment of a sample collection device (60) in accordance with this invention which is particularly intended for
10 use in immunodiagnostic testing for faecal occult blood, the device comprising a dipstick collection device (61) and a protective cover member or housing (62) adapted to receive and shield the dipstick collection device (61). As shown in Figure 6B, an immunochromatographic test strip may be adhered to one side of the protective cover member (62) and used in the immunodiagnostic test by
15 extraction of analyte from the absorbent matrix of dipstick collection device (61) by addition of an extraction reagent to the absorbent matrix (63) thereof, and then transferring the extraction reagent from the dipstick device (61) to the immunochromatographic test strip (64) as described in detail below.

Figures 7A, 7B and 7C illustrate an alternative embodiment of the protective cover member in accordance with this invention, with this embodiment being formed by folding a single piece of cardboard as shown in Figure 7A to give a two-panel protective cover member (71). Member (71) is provided with apertures (72) and (73) in the panels thereof with aperture (72) being located so
25 as to correspond with at least a portion of the absorbent matrix (77) of the dipstick collection device (74) when device (74) is assembled within panel A of the protective cover member (71). In use of the device shown in these figures, an immunochromatographic test strip (75) is located on panel B of the protective cover member (71) as illustrated in Figure 7C, and extraction reagent is added
30 to the reagent pad (76) at one end of the test strip and the protective cover member (71) closed. As described below, the extraction reagent flows from the reagent pad (76) along test strip (75) via the specimen matrix (77) of the dipstick

- 16 -

(74) which it contacts through window (72) and then to the remaining components of the test strip, as is already well known. Since the test strip (75) is provided with a clear plastic backing, the result of the test can be observed through the window (73) in panel B.

5

The modified dipstick collection device (81) shown in Figures 8A and 8B comprises a semi-rigid elongate backing (82) having a sample collection matrix (83) adhered towards one end thereof, and in operable contact therewith an absorbent reagent matrix (84). An impermeable protective membrane layer (85) is provided which covers at least the junction between the sample collection matrix (83) and the absorbent reagent matrix (84) to prevent contamination of matrix (84) while still allowing test reagent(s) added to matrix (84) to flow along the device (81) and into the sample collection matrix (83).

15 As an alternative, absorbent reagent matrix (84) may be replaced by capillary means to conduct test reagent(s) to the sample collection matrix (83).

The protective cover member or sheath (86) shown in Figure 8C is provided with a port or window (87) through which test reagent(s) may be applied to the absorbent reagent matrix (84) when the device (81) is received within the sheath (86), together with a transparent reading window (88) which registers with the sample collection matrix (83). Note that part of the sample collection matrix (83) may be covered by the protective membrane layer (85) to provide a clear region of the sample collection matrix for observing development of colour on the test paper.

Figures 9A and 9B show a modified version of the sample collection device of Figure 7, with the two-panel protective cover member (91) being formed by folding a single piece of cardboard, as shown in Figure 9A. As shown in Figure 9B, the sample collection device used in this modified version may be a modified dipstick device (81) as shown in Figures 8A and 8B, and described above.

- 17 -

In Figure 9B, the right-hand panel of protective cover member (91) is provided with a recess (92) for insertion of dipstick device (81), and recess (92) is covered with a protective cover layer (93) which covers most of the dipstick device (81) when it is located within the cover member (91), but at least a portion of the sample collection matrix (83) is left exposed. A reagent port (94) is provided in the cover layer (93) for addition of test reagent(s) to the absorbent reagent matrix (84) of dipstick device (81). Window (95) is provided in the left-hand panel of cover member (91) to register with an immunochromatographic test strip (96). If desired, the test strip (96) may be mounted on a test strip insert (97) and a locating recess (98) provided in the left-hand panel of cover member (81) to ensure that, in use, test strip (96) is properly located with respect to both the dipstick device (81) and the window (95) through which the result of the test is to be observed.

The following detailed description relates to sample collection devices according to the present invention which are intended for faecal occult blood detection, both by guaiac and immunodiagnostic procedures. The collection devices of the invention have been developed with minimum-cost manufacture as the foremost consideration, but with ease and convenience of use as important secondary considerations. The devices are designed for manufacture by established high speed, fully automated commercial procedures, for example by web handling technology, and provide the basis for simple and inexpensive test kits.

Both collection devices use an elongate collection member referred to hereafter as a dipstick, as shown in Figures 1A and 1B. An absorbent matrix such as absorbent paper is adhered to one side of one end of a semi-rigid backing such as polycarbonate, Mylar, high impact polystyrene or waterproofed cardboard. In use, the dipstick is inserted into the faecal material and smeared to ensure complete and even penetration of the absorbent matrix by the specimen. Excess faecal material is then removed from the dipstick by one of several possible means. For example:

- 18 -

- (i) the dipstick may simply be wiped clean by a suitable, readily disposable absorbent material such as toilet tissue;
- (ii) plastic backed absorbent paper, such as used for covering laboratory benches, may be folded into a simple sleeve and used to wipe the dipstick clean as the dipstick is withdrawn from the sleeve. Alternatively, a pre-formed sleeve may be provided ready for use, or already in place around the stem of the dipstick (Figure 2);
- (iii) the dipstick may be supplied in a disposable, protective sheath which may be discarded with excess faecal material after sample collection (Figure 10); or
- (iv) the dipstick may be manufactured with a peel-off impermeable cover (with or without a filter screen) as shown in Figures 4A and 4B, and after faecal collection, the cover is peeled-off and discarded.

Guaiac tests require dietary restrictions for several days before testing in order to avoid possible false positive results that may be caused by a number of foodstuffs and drugs. This limits the appeal and reliance that may be placed on this otherwise simple test. Immuno-based tests give specificity for human blood, but are more expensive and complex to perform.

These difficulties may be overcome by firstly performing mass screening using the guaiac test, but without dietary restrictions, and subsequently confirming guaiac positive specimens using an immunochromatographic test.

The "dipstick and sheath" approach of the present invention is ideal for this testing strategy, as one side of the dipstick may be provided with guaiac paper, the other with collection paper for an immuno-based test, both according to one of the several variations described herein. Alternatively, two dipsticks could be laminated together to achieve the same effect. Faecal specimen

- 19 -

samples would thus be automatically collected simultaneously for both test types.

5 Upon identification of a guaiac-positive specimen, the dipstick containing the specimen would be removed from the sheath, turned over and the result confirmed by an immuno-test. This allows confirmation of the result on the same faecal specimen. As bleeding in many cases may be intermittent, performing confirmatory testing on later specimens may not be as useful.

10 1. **Guaiac testing.**

Guaiac testing is the most widespread technology for faecal occult blood testing. The tests are rapid, inexpensive and easy to use, but lack specificity for human haemoglobin. Typically (e.g. SmithKline Diagnostics' Hemoccult), a
15 faecal specimen is smeared on a test paper inside a test card and sent to the physician's office or laboratory for test development. Developer solution is added to the reverse side of the paper in the card and the development of a blue colour in the paper indicates a positive detection of haemoglobin. Some food materials (e.g. horseradish or haemoglobin from food animals) may cause
20 false positive results with guaiac tests.

In using the collection device of this invention with guaiac procedures, the dipstick serves as both a collection and testing member, with the absorbent matrix being guaiac test paper. After the sample has been collected as
25 described above, the test may be developed by the addition of developer solution to the test (guaiac) paper, and the result (blue colour development) read directly by observing the test paper or, if the dipstick has a transparent plastic carrier, indirectly through the carrier.

30 If desired, the dipstick may be provided with a covering membrane as shown in Figures 3A and 3B which is laminated to the semi-rigid backing in such a way that a portion of the guaiac test paper making up the absorbent matrix is

- 20 -

protected from contact with the faecal sample. In use of such a device, developer solution added to the guaiac test paper is allowed to chromatograph away from the faecal sample and into the protected portion of the test paper, allowing development against a clear background and hence a clear region for
5 observing the development of colour on the test paper.

In many cases, the collection of a sample may be done by a patient at home and the device handed to a physician and then sent to a laboratory for completion of the testing. In these cases, the wiped dipstick would be inserted
10 into the paper, cardboard or plastic sheath or other protective cover member on which the patient's name, and possibly test instructions, could be written. Testing at the laboratory could then be completed by simply removing the dipstick from the sheath, adding developer solution and reading as described above.

15

Alternatively, the sheath may be constructed as shown in Figures 5A and 5B in order to protect the patient, physician and test operator from exposure to the faecal sample. In this case, the dipstick would have a clear plastic carrier backing and would be placed in the sheath or other protective cover member so
20 that the guaiac test paper was located below an aperture in the sheath. The operator would add a drop of developer solution to the test paper via the aperture, and turn the sheath over to read the result via the reading window in the other side of the sheath. (For additional containment, the aperture may be covered with a resealable, peel-back sealing layer and, if desired, the window
25 sealed with a clear sealing layer.)

In the modification shown in Figures 8A and 8B, the dipstick has been modified by the addition of an absorbent reagent matrix above, and in operable contact with, the sample collection matrix, such as guaiac faecal collection
30 paper. The flow of test reagent is therefore changed along the dipstick. The region of the junction between the sample collection matrix and the reagent matrix is preferably covered with a protective membrane, which serves to protect

- 21 -

the reagent matrix from faecal contamination. The protective membrane may also be printed to indicate the depth limit for faecal smearing, and the orientation of the dipstick for its insertion in the sheath.

- 5 One advantage offered by these modifications is that the absorbent matrix at the top of the sample collection matrix assists rapid drying of a faecal specimen. The moisture is drawn from the sample collection matrix by the absorbent and vented via the reagent port in the sheath. Rapid drying of the faecal specimen is important in order to avoid degradation of the analyte. The
- 10 absorbent matrix may also be pre-impregnated with materials for removing or disguising any odours from the faecal specimen.

 Addition of the test reagent to the absorbent matrix also ensures that its flow is controlled and that there is an orderly and even application of reagent to

15 the top of the sample collection matrix. Furthermore, for guaiac testing, containment is improved as only the clean absorbent reagent matrix is exposed at the reagent addition port.

 In this modified embodiment, the dipstick collection device may have a

20 band of guaiac paper, with its lower region protected from faecal contact by a clear protective membrane. A band of absorbent material (e.g. Ahlstrom 1281 or 939 paper) is in operable contact with the upper margin of the guaiac paper. The upper margin of the guaiac paper and the lower region of the absorbent are covered with an adhesive membrane, which serves to protect the absorbent from

25 faecal contact. Alternatively, the absorbent material may be replaced by a capillary device.

 The protective cover member or sheath is also modified to allow reagent addition directly to the absorbent paper, which acts as a reservoir for orderly and

30 even application of the developer to the guaiac paper. Preferably the sheath would be formed of a clear plastic material, so that the downward migration of

the developer solution may be visualised in the guaiac paper, without exposure to the faecal-contaminated surface of the guaiac paper.

The sample collection device described above enables simple faecal occult blood testing kits to be made available particularly to developing countries and cost-sensitive markets. Such test kits may consist of bulk packaged dipsticks and sheaths, together with an appropriate supply of developer solution. Such test kits would significantly reduce the cost of guaiac testing for faecal occult blood since the dipstick uses minimal and inexpensive materials relative to current test systems, the dipstick may be manufactured by a continuous, high speed, low cost manufacturing process, whereas existing tests involve more manufacturing stages and greater complexity; insertion of the dipstick into the specimen reduces exposure to the specimen and avoids the requirement to include a smearing stick or paddle in the kit, and the small number of kit components and compact nature of the components ensures low packaging, transport and storage costs.

2. Immunodiagnostic testing.

Immunodiagnostic testing allows absolute specificity for human haemoglobin in faecal occult blood testing (e.g. SmithKline Diagnostics' HemeSelect), but entails some increase in test cost and complexity. In some markets, immunodiagnostic tests will be essential in order to avoid interference from commonly used foodstuffs. If available at a sufficiently low cost, these tests could ultimately replace guaiac tests.

For immunodiagnostic testing using the sample collection device of this invention, the dipstick may act simply as a collection instrument for subsequent transfer of faecal extract to an immunodiagnostic test device, or the dipstick may be used both for sample collection and, in conjunction with a protective sheath, as a test device. The absorbent matrix on the dipstick is a suitable collection paper (e.g. Whatman's #1) and the dipstick is inserted into the faecal material,

- 23 -

removed, cleaned and sheathed in the protective cover member as already described. The faecal collection matrix may then be extracted immediately to yield a test sample, or sent to the physician's office or laboratory for subsequent extraction and completion of testing. There the dipstick may be unsheathed and
5 the faecal material extracted from the collection matrix by the addition of one or more drops of extraction reagent, typically a buffered detergent solution such as PBS-Tween 20.

For example, extraction reagent may be added to the absorbent matrix
10 with the collection device in the horizontal position. The reagent remains in contact with the matrix thereby allowing mobilisation of any haemoglobin into the reagent. The liquid extract may then be transferred from the dipstick to an immunodiagnostic test device by simply elevating the handle end of the dipstick and pouring the drop from the dipstick to the test device. The extract may be
15 transferred to a microwell, tube or membrane for EIA testing, or to the sample port/specimen pad of an immunochromatographic device for "one-step" diagnostic testing. A suitable immunochromatographic device for diagnostic testing is described in Australian Patent Application No. 21852/92 (International Patent Application No. PCT/US92/04425) referred to above, the contents of
20 which are incorporated herein by reference.

A simple low-cost diagnostic kit, using the collection device of this invention and intended particularly for "cost-conscious" applications, is illustrated in Figures 6A, 6B and 6C.
25

The faecal specimen is collected on the absorbent matrix and the dipstick inserted into a sheath that has printed on it the patient's name, a marked position indicating the placement for an immunochromatographic test strip and the instructions for the test. The immunochromatographic test strips are
30 packaged in bulk in a resealable container with sufficient desiccant to maintain dry conditions. A strip is taken from the container, protective paper removed from its underside to expose adhesive and the strip affixed via the adhesive to

- 24 -

the position outlined on the sheath. The end of the dipstick where the absorbent matrix is located is placed in a horizontal position in contact with the sample receiving pad of the test strip. Extraction reagent is added to the absorbent matrix and the dipstick tipped to transfer the specimen extract to the test strip.

- 5 With most immunochromatographic devices, the results typically develop in the test strip within five minutes.

A marginally more complex version of the kit, which offers improved containment of any pathogenic organisms in the sample, is shown in Figures 7A,
10 7B, 7C and 7D.

The sheath is formed from a single piece of cardboard, folded as shown in Figure 7A to give a two-panel housing. The housing is shown in "open" form in Figure 7B and 7C and may be closed by folding the panels together prior to
15 use.

The closed (spot glued) housing is issued to the patient who collects the specimen and inserts the dipstick into panel A, which acts as the sheath. The physician opens the housing and adheres a test strip to the position indicated
20 in dotted outline on panel B. The extraction reagent is added to the reagent pad at the base of the test strip and the housing sealed closed. The liquid flowpath is as shown diagrammatically in Figure 7D. The extraction reagent flows from the reagent pad, via the specimen matrix of the dipstick (thereby mobilising any haemoglobin), to the remaining components of the test strip. The test strip is on
25 a clear plastic backing, which allows the result to be observed through the window of the closed housing.

This test format offers maximum containment and ease of use, and suitable devices for immunochromatographic testing utilising this test format are
30 described in Australian Patent Application No. 21852/92, above.

- 25 -

Figures 9A and 9B illustrate a modified version of the device of Figures 7A to 7D. In most circumstances it is expected that the patients will collect their own faecal specimens using the dipstick, which will be returned, within its sheath, to a health professional for test development. As previously indicated
5 in the description of Figure 7, the sheath will preferably also be the test housing.

A modified version of the Figure 7 test format, suitable for most markets, is shown in Figure 9B. As distinct from the format illustrated in Figure 7, in this modified version the test reagent is added to a reagent pad situated on the
10 dipstick, instead of on the test strip. It should be noted that in this modified version, the amount of faecal specimen tested, and hence the test sensitivity, is controlled by the gap width between the upper and lower protective membranes. This can be set and controlled during manufacture for optimal test performance. This format of the device is intended for faecal specimen smearing by the patient
15 and subsequent test development by the health professional. In this case, the dipstick would be smeared by the patient and inserted in the test housing for return for test development. For this format, a test kit would comprise: one test housing and dipstick, and one transport envelope (optional), issued to the patient; and bulk packaged test strips (desiccated), and a dropper bottle of
20 buffer, issued to the laboratory/health professional for test development.

In the test method using this format, the patient smears the dipstick with faecal specimen, and then inserts the wiped dipstick into the housing (sheath) and returns them (in the transport envelope) for development. The
25 laboratory/health professional opens the test housing, inserts the keyed test strip in the recess of the left hand panel, adds the specified number of drops of reagent to the reagent pad of the dipstick, and then seals the test housing closed and reads the result through the window in the standard manner.

30 For situations where the smearing of the dipstick and test development is done at the same site, e.g. by the laboratory/health professional or by the test

subject for home testing, the test housing may be manufactured and issued with the test strip in place.

In this format, a test kit would consist of a test housing, issued in a desiccated pouch, and a dipstick. The test procedure would require the operator to smear and wipe the dipstick and insert it into the test housing. The test would be initiated by addition of reagent to the dipstick as already described.

In some markets, e.g. in Japan, the patient may be unwilling to smear his or her faecal specimen on the dipstick collection device, preferring instead to present the faecal specimen directly to the laboratory or health professional for testing. In addition, the Japanese diet causes many false positives with guaiac testing, making specific immunodiagnostic testing the only viable option.

A number of different formats of the device of the present invention are particularly suitable for immunodiagnostic testing in this market.

Firstly, a sheathed dipstick collection device of the type illustrated in Figure 10 may be used as it is quite inexpensive and therefore suitable for mass-screening programs. In use, such a collection device is dipped into the faecal specimen so that faecal material contacts the absorbent matrix through the opening in the sheath, and the dipstick collection device is then removed with the sheath being discarded into the specimen container. The dipstick device having the sample applied can then be inserted into a protective cover member provided with one or more ports or openings for addition of test reagent, as well as one or more reading windows, so that the appropriate test reagent can be added and the test result read with the dipstick device *in situ* within the protective cover member. (Alternatively, the protective cover member could have the test reagent sealed in the base and the dipstick device could, on insertion, perforate a frangible seal to automatically release the reagent and initiate test development.) In another option, the dipstick device having the sample applied may be inserted into a transparent tube or the like as a

- 27 -

protective cover member with test reagent already present or being added to the tube prior to insertion of the dipstick device.

Alternatively, a hybrid guaiac/immunodiagnostic test is also possible. In this hybrid test, a guaiac dipstick collection device, as already described, would have a band or spot of antibody specific for human haemoglobin immobilised within the guaiac paper. Typically, the antibody would be coated on latex beads of a size sufficient to prevent their migration within the guaiac paper. Addition of developer to one margin of the guaiac paper would cause migration of collected sample across the paper and the specific accumulation of human haemoglobin at the antibody band or spot. The guaiac developer would indicate this accumulation by a concentrated colour development in the region of the antibody. The use of an antibody in such a hybrid test would require that the dipstick should be issued in a desiccated package (to protect the antibody), and that the test should be developed soon after faecal addition, in order to prevent degradation of the antibody.

3. Combined Guaiac/Immunodiagnostic testing.

For cost-effective, mass-screening program for detection of faecal occult blood in at-risk populations (e.g. 50 years and older), a combined guaiac/immunodiagnostic test is also feasible which allows guaiac positives to be confirmed by an immunodiagnostic format, (see Allison *et al.*, 1996, *supra*). In such a combined test, for example using the device described in Figures 9A and 9B, the dipstick collection device may be provided with a guaiac absorbent matrix on one side of the elongate carrier and a second absorbent matrix on the opposite side of the carrier for collection of sample material for immunodiagnostic testing. The protective cover member provides a sheath or housing for transport of the dipstick collection device, and in addition may be configured as previously described to provide a test housing with ports or windows for addition of test reagent and reading of test results in the guaiac test. In use, the dipstick collection device is dipped into a faecal sample thereby

- 28 -

collecting sample material on both sides, and excess material is then wiped off prior to insertion of the collection device into the protective cover member.

Testing of the collected samples by the laboratory or health professional
5 may then simply comprise carrying out of guaiac testing on one side of the dipstick device *in situ* in the protective cover member as a test housing, and where confirmation of a guaiac positive result is required, elution of the specimen from the absorbent matrix on the other side of the device by addition of buffer, with the eluted specimen being transferred directly onto an
10 immunochromatographic test strip as described in Australian Patent Application No. 21852/92, above.

Such a combination guaiac/immunodiagnostic test offers significant advantages in that it allows two types of tests on the same faecal specimen, and
15 thus enables direct immunodiagnostic confirmation of a guaiac-positive test result. In particular, it provides a system for cost-effective mass screening of at risk populations in which a guaiac screen can be carried out without dietary restriction and as described above guaiac-positive results can be confirmed by an immunodiagnostic procedure on the same specimen.

20

With test kits as described above, immunochromatographic testing may be rendered sufficiently inexpensive as to be feasible for developing countries, as the dipstick collection device may be mass produced using very high speed,
25 low-cost manufacturing procedures; the sheath serves both for transfer of the dipstick collection device to the test site and also as the base and instruction card for the test itself (this rationalises material usage and kit complexity); the test strips, through bulk packaging in a desiccated container, avoid the usual expensive requirement for individual desiccated packaging in a foil pouch; and
30 the minimal size and number of test components ensures the inexpensive packaging, storage and transport costs.

CLAIMS:

1. A sample collection device comprising:
 - (a) a collection member having at least one absorbent matrix thereon, and
 - (b) a protective cover member for said collection member, said protective cover member being adapted to receive and shield at least the or each absorbent matrix of said collection member when the collection member is assembled therewith.
2. A device according to claim 1 wherein said collection member comprises an elongate member in the form of a dipstick, test strip or the like.
3. A device according to claim 2 wherein said elongate member comprises an elongate rigid or semi-rigid carrier or backing having the or each absorbent matrix affixed to the carrier or backing at or towards one end thereof.
4. A device according to claim 3 wherein said carrier or backing comprises a material selected from plastics materials, water-proofed or water-resistant cardboard, or the like.
5. A device according to claim 4 wherein said carrier or backing comprises a transparent or translucent material.
6. A device according to any of claims 1 to 5 wherein the or each absorbent matrix comprises an absorbent material.
7. A device according to claim 6 wherein said absorbent material is selected from filter paper, cellulose, nitrocellulose, nylon, rayon, glass fibre, sintered glass, fleece and non-woven or porous synthetic materials.

8. A device according to any of claims 1 to 7 wherein the or each absorbent matrix comprises a sample collection matrix in operable contact with an absorbent reagent matrix or capillary means, wherein reagent added to the absorbent reagent matrix or capillary means can flow along the collection member and into the sample collection matrix.
9. A device according to claim 8 wherein at least the region of the junction between said sample collection matrix and said absorbent reagent matrix or capillary means is covered with an impervious protective layer to prevent contamination of the absorbent reagent matrix or capillary means when a sample is applied to the sample collection matrix.
10. A device according to any of claims 1 to 9 wherein said collection member is provided with a protective cover layer or sheath.
11. A device according to claim 10 wherein said protective cover layer or sheath is provided with a window or aperture exposing the or each absorbent matrix or a portion thereof.
12. A device according to claim 11 wherein a filter material is located over the exposed portion of the or each absorbent matrix.
13. A device according to any of claims 10 to 12 wherein said protective cover layer or sheath is removable.
14. A device according to any of claims 1 to 13 wherein said protective cover member comprises a sheath or housing having an internal recess adapted to receive said collection member and shield the or each absorbent matrix thereon.

- 31 -

15. A device according to claim 14 wherein the internal recess in said sheath or housing receives substantially the entire length of an elongate collection member.
16. A device according to claim 14 or claim 15 wherein said protective cover member comprises a water-impervious non-absorbent material selected from plastics materials, water-proofed or water-resistant cardboard or similar materials, and laminated materials.
17. A device according to any of claims 14 to 16 wherein said protective cover member is provided with at least one window or aperture through a wall or panel thereof, the or each window or aperture being located so that it corresponds with at least a portion of an absorbent matrix when the collection member is assembled within the protective cover member.
18. A device according to claim 17 further comprising a removable, and optionally resealable, sealing layer covering the or each window or aperture in the protective cover member.
19. A device according to claim 17 or claim 18 wherein the protective cover member is provided with at least one additional window or aperture through a wall or panel thereof, said additional window or aperture located so that, in conjunction with the use of a transparent or translucent carrier for the collection member, a reading window or aperture is provided for reading the results of a test through the transparent or translucent carrier.
20. A device according to any of claims 14 to 19 wherein the protective cover member includes test reagent(s) sealed therein with a frangible seal, whereby said frangible seal may be broken or perforated when said collection member is inserted into the protective cover member.

- 32 -

21. A device according to any of claims 14 to 20 wherein said protective cover member comprises:
 - a first opposable component adapted to receive said collection member, and
 - a second opposable component,wherein said first and second opposable component can be brought into opposition.
22. A device according to any of claims 1 to 21 wherein at least a portion of the or each absorbent matrix is impregnated with guaiac.
23. A device according to any of claims 1 to 22 wherein said collection member comprises two or more absorbent matrices, and at least a portion of one of said matrices is impregnated with guaiac.
24. A device according to any of claims 1 to 23 further comprising a test strip comprising means for detecting an analyte in a sample, said test strip being located or adapted to be located on said collection member or on or in said protective cover member.
25. An assay device for detection and/or determination of an analyte in a sample, comprising:
 - (a) a collection member having at least one absorbent matrix thereon adapted to receive said sample,
 - (b) a protective cover member for said collection member, said protective cover member being adapted to receive and shield at least the or each absorbent matrix of said collection member when the collection member is assembled therewith, and
 - (c) means to detect and/or determine said analyte in said sample.

- 33 -

26. An assay device according to claim 25, wherein said means to detect and/or determine said analyte comprises guaiac impregnated in at least a portion of the or each absorbent matrix.
27. An assay device according to claim 25, wherein said means to detect and/or determine said analyte comprises a test strip for detecting said analyte in said sample.
28. An assay device according to claim 27 wherein said test strip includes a chromatographic medium.
29. An assay device according to claim 27 or claim 28, wherein said test strip is located or adapted to be located on said collection member.
30. An assay device according to claim 27 or claim 28 wherein said test strip is located or adapted to be located on or in said protective cover member.
31. An assay device according to claim 30 wherein said protective cover member comprises:
 - a first opposable component adapted to receive said collection member, and
 - a second opposable component,said test strip is located or adapted to be located on or in said second opposable component, and said first and second opposable components can be brought into opposition so as to cause the sample to be brought into contact with said test strip.

1/7

FIG 1A



FIG 1B

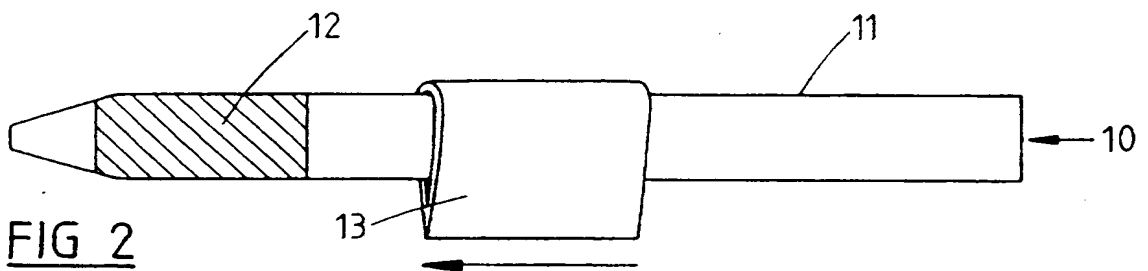


FIG 2

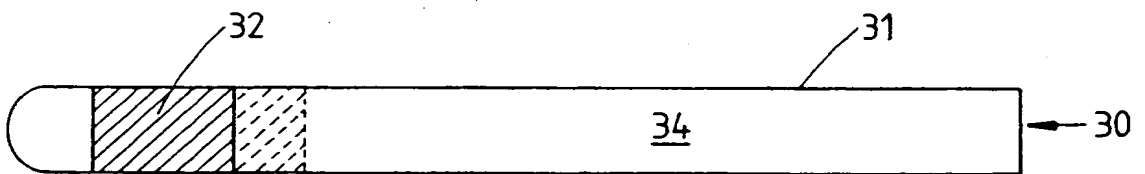


FIG 3A

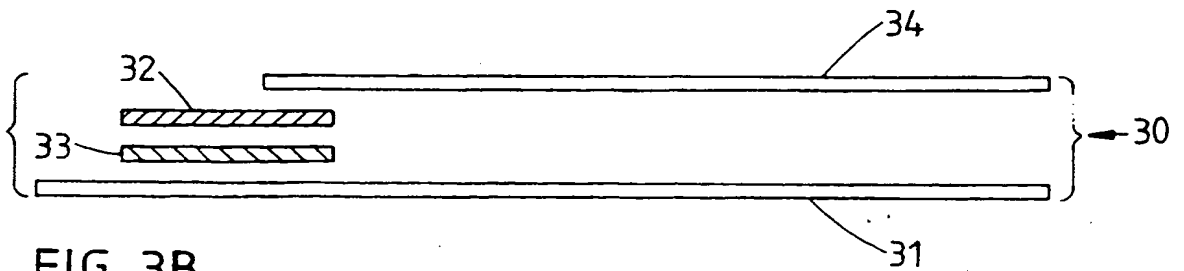
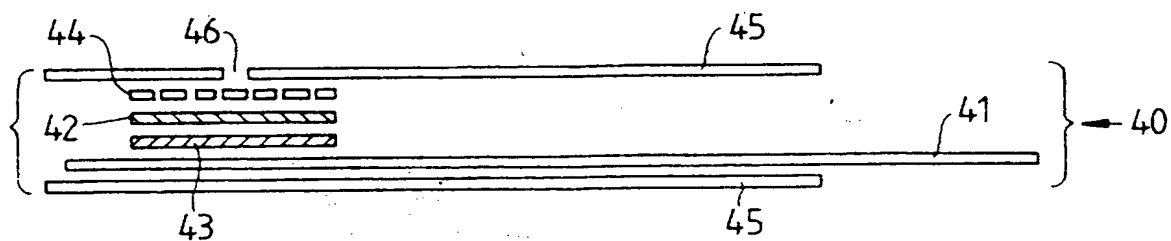
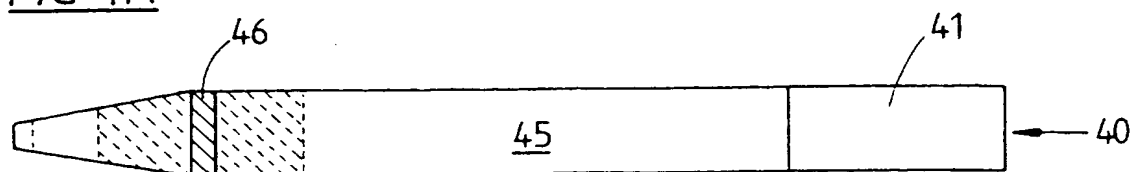
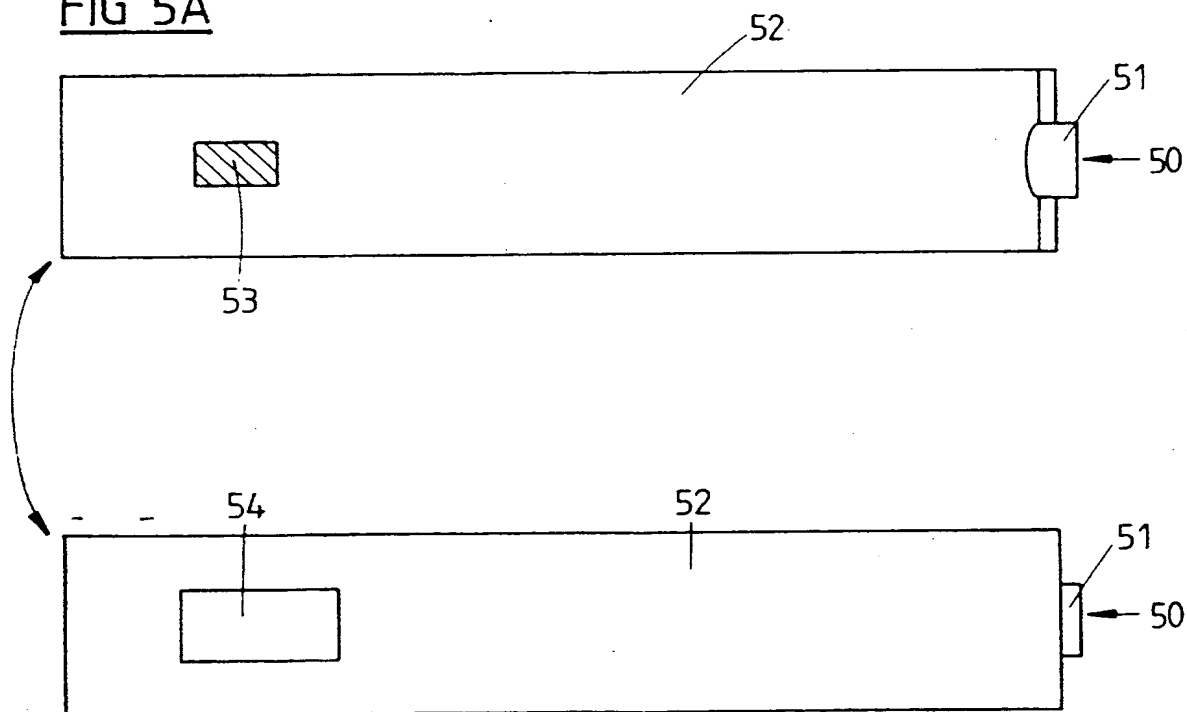
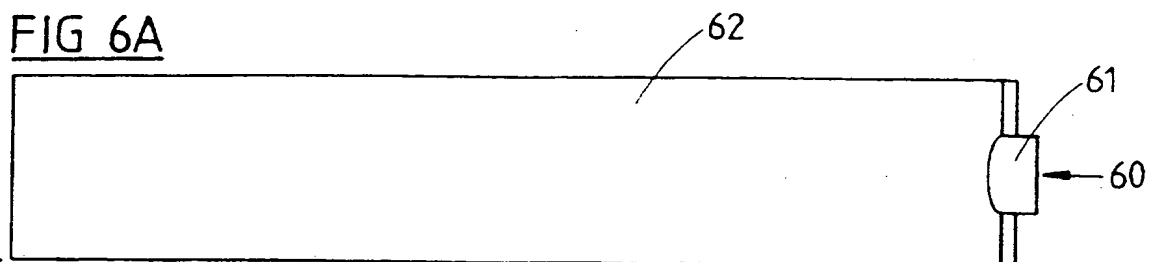
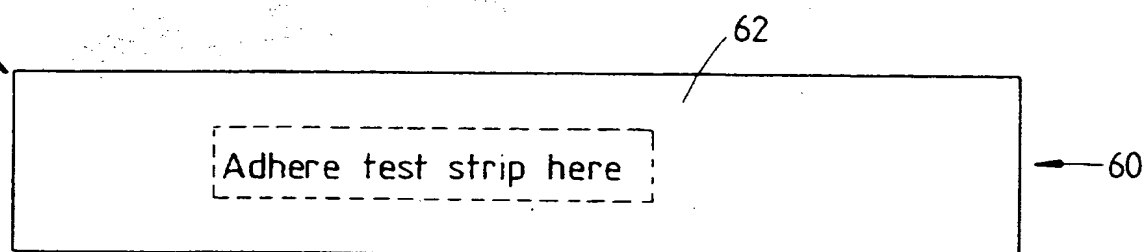
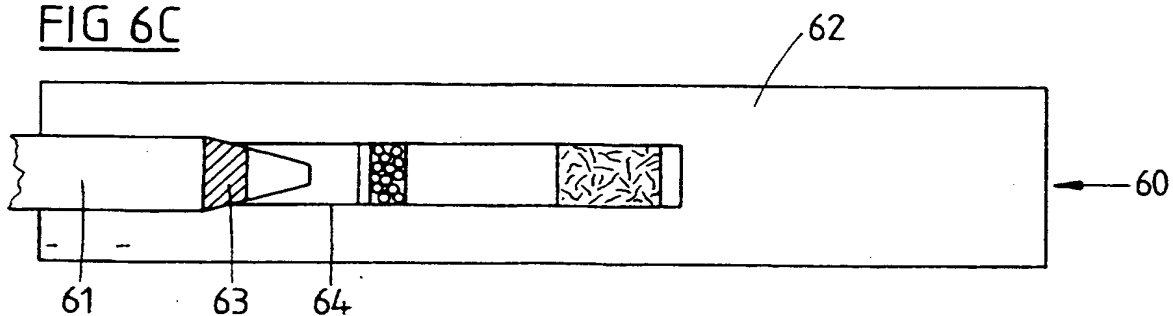


FIG 3B

2/7

FIG 4AFIG 4BFIG 5AFIG 5B

3/7

FIG 6AFIG 6BFIG 6C

4/7

FIG 7A

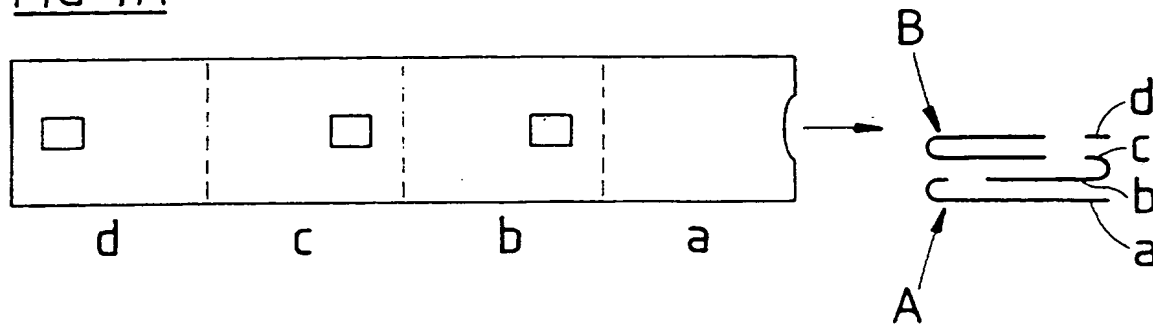


FIG 7B

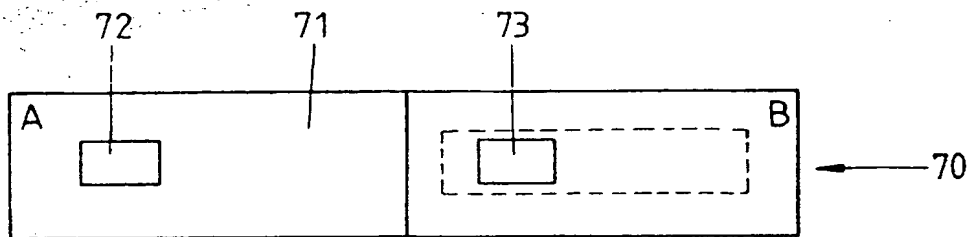


FIG 7C

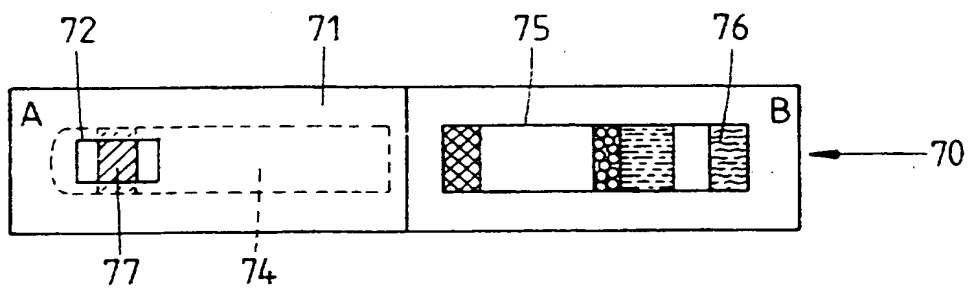
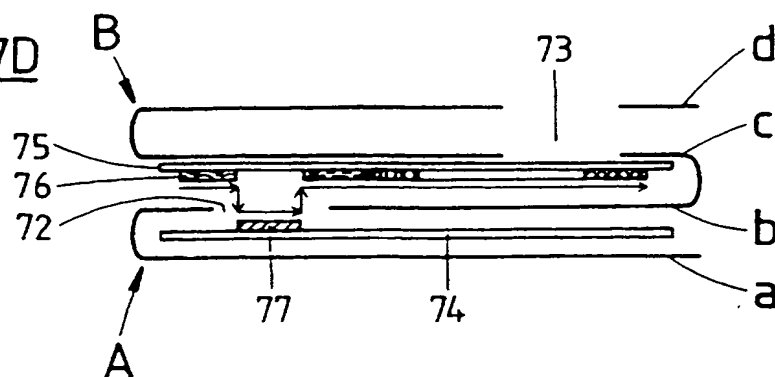
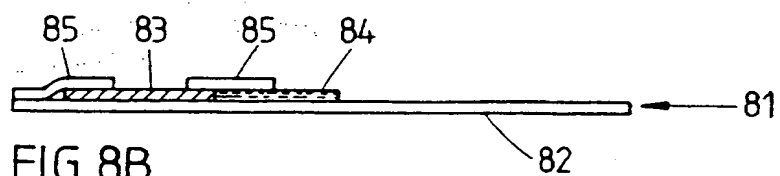
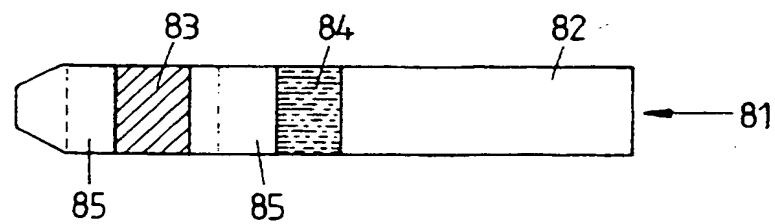
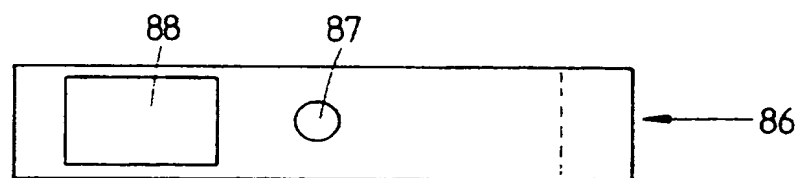
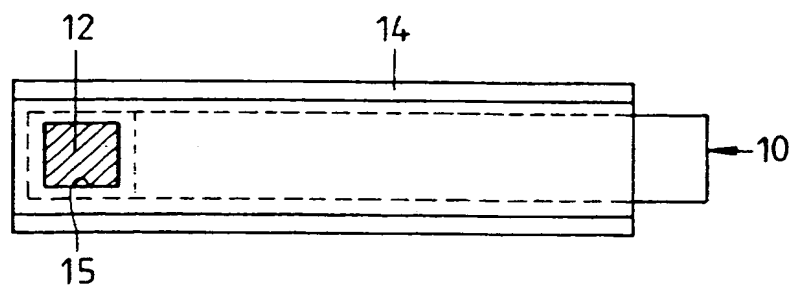


FIG 7D



5/7

FIG 8AFIG 8BFIG 8CFIG 10

6/7

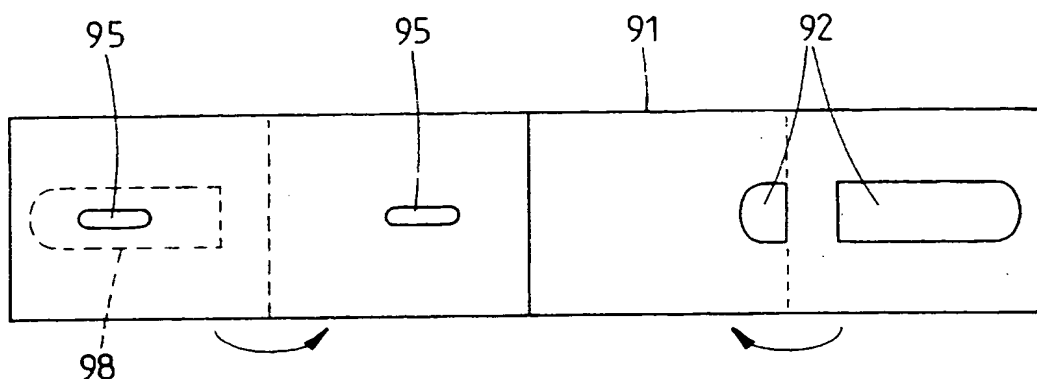


FIG 9A

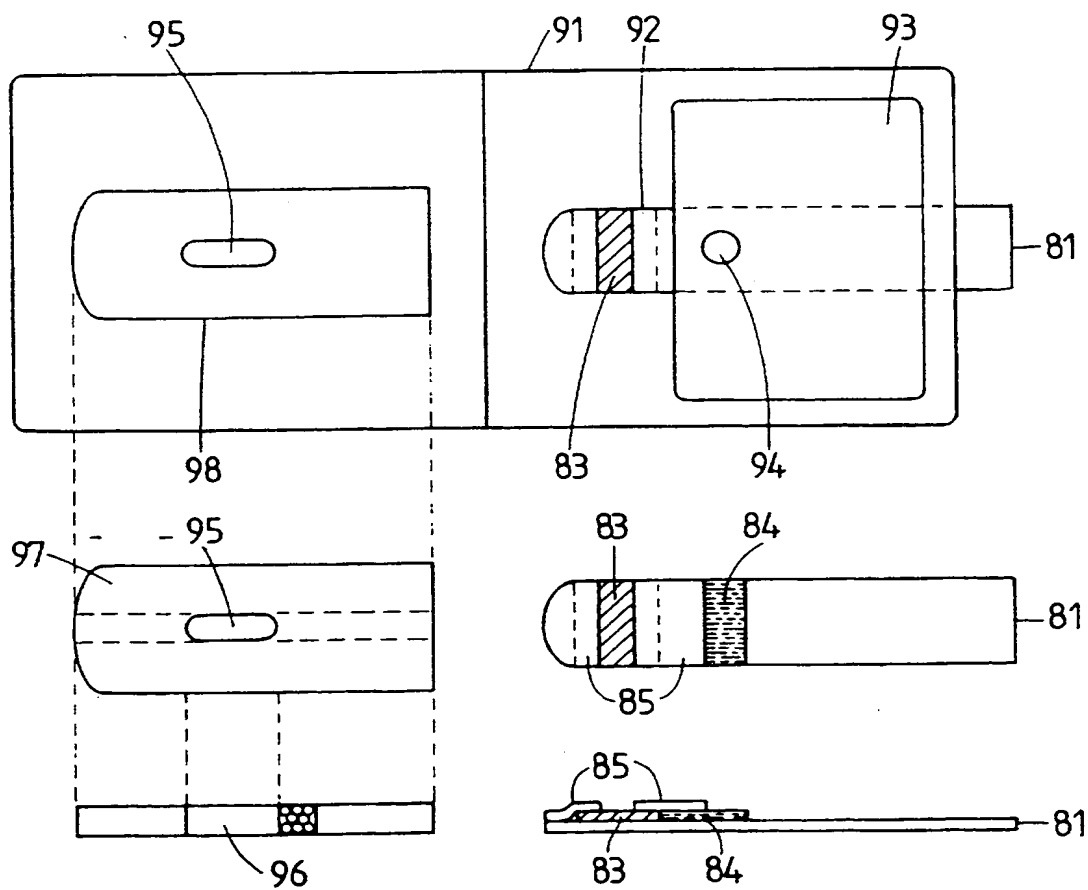
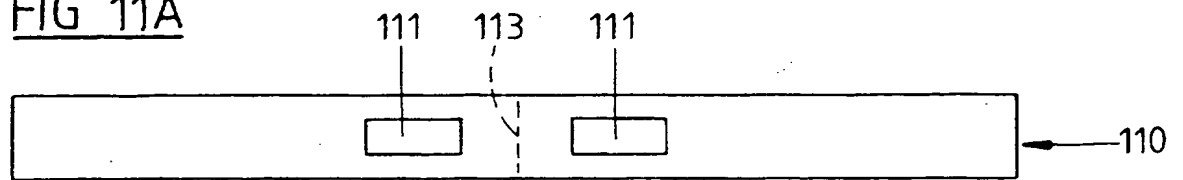
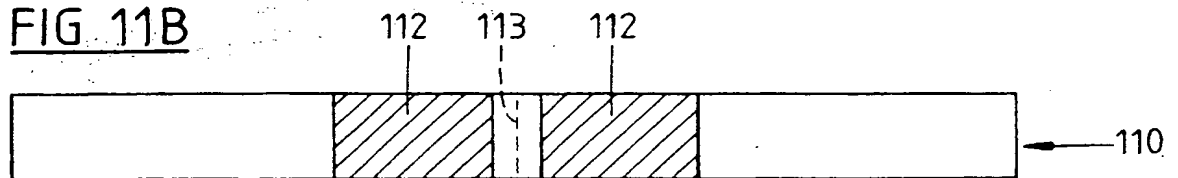
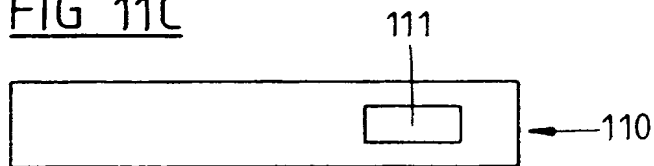
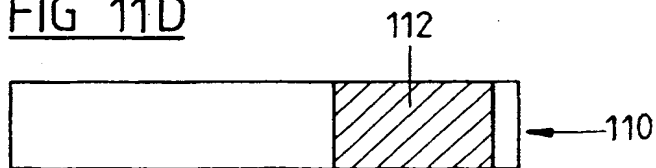


FIG 9B

7/7

FIG 11AFIG 11BFIG 11CFIG 11D

INTERNATIONAL SEARCH REPORT

International Application No.
PCT/AU 96/00135

A. CLASSIFICATION OF SUBJECT MATTER		
Int Cl ^o : G01N 1/00, 1/28, 30/90, 33/53		
According to International Patent Classification (IPC) or to both national classification and IPC		
B. FIELDS SEARCHED		
Minimum documentation searched (classification system followed by classification symbols) IPC: G01N 1/00, 1/02, 1/10, 1/28, 30/90, 30/91, 31/08, 33/53, 33/54, 33/543, 33/545, 33/55, 33/551, 33/552		
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched AU: IPC as above		
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) JAPIO, WPAT		
C. DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	EP279574 A (METAL BOX PLC) 24 August 1988 Column 2 line 31 to column 4 line 10	1-4, 6-11, 14-20, 24, 25, 27-30
Y		12, 22, 23, 26
P.X	AU 79941/94 A (UNIPATH LTD) 18 May 1995 Page 7 line 5 to page 9 line 27, figs.	1-4, 6-10, 13, 14, 16, 17, 25, 27, 28, 30, 31
P.Y		12, 22, 23, 26
<input checked="" type="checkbox"/> Further documents are listed in the continuation of Box C <input checked="" type="checkbox"/> See patent family annex		
* Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "&" document member of the same patent family		
Date of the actual completion of the international search 2 May 1996		Date of mailing of the international search report 10TH MAY 1996
Name and mailing address of the ISA/AU AUSTRALIAN INDUSTRIAL PROPERTY ORGANISATION PO BOX 200 WODEN, ACT 2606 AUSTRALIA Facsimile No.: (06) 285 3929		Authorized officer S. CLARK Telephone No.: (06) 283 2164

INTERNATIONAL SEARCH REPORT

Int. National Application No.

PCT/AU 96/00135

C (Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	EP 306336 A (SYNTEX (U.S.A.) INC.) 8 March 1989 Column 13 line 37 to column 15 line 52	1-4, 6-11, 13-17, 19, 21, 24, 25, 27-31
Y		12, 22, 23, 26
X	EP 253579 A (HYBRITECH INC.) 20 January 1988 Column 6 line 47 to column 7 line 48	1, 6-10, 12, 13, 25
Y		22, 23, 26
X	EP 323605 A (ABBOTT LABS) 12 July 1989 Page 10 line 38 to page 11 line 5, figs.	1-9, 24, 25, 27-31
Y		12, 22, 23, 26
P,X	AU 12659/95 A (SMITHKLINE DIAGNOSTICS INC.) 15 June 1995 See pages 38, 39	1-4, 6-11, 24, 25, 27-30
P,Y		12, 22, 23, 26
X	US 5064766 A (WARDLOW) 12 November 1991 Column 4 line 4 to column 6 line 42	1, 6-11, 13, 22, 23, 25-28
Y		12
Y	EP 269362 A (SMITHKLINE DIAGNOSTICS INC.) 1 June 1988 Page 3 line 50 to page 4 line 44	22, 23, 26

International Application No.
PCT/AU 96/00135

Information on patent family members

This Annex lists the known "A" publication level patent family members relating to the patent documents cited in the above-mentioned international search report. The Australian Patent Office is in no way liable for these particulars which are merely given for the purpose of information.

Patent Document Cited in Search Report				Patent Family Member			
EP	279574	AU	11458/88	BR	8800577	CA	1318580
		DK	795/88	FI	880698	GB	2201241
		HK	316/91	JP	63223559	NO	880665
		SG	45/91	US	4965047	ZA	8800906
AU	79941/94	WO	9513541	EP	653639	FR	2712392
		US	5504013				
EP	306336	CA	1333046	JP	1072066	US	4981786
US	5064766	US	5171528				
EP	323605	AU	27095/88	CA	1312544	CN	1036078
		JP	1244370				
AU	12659/95	WO	9516207				
EP	253579	AU	75571/87	CA	1288690	JP	63042471
		US	5120504				
EP	269362	AU	81636/87	CA	1309330	DK	6037/87
		FI	875173	JP	63142262	NO	874868
		PT	86201	US	4789629	ZA	8708761
END OF ANNEX							

This Page Blank (uspto)